

Applicant : Tamar H. Michaeli  
Serial No. : Unknown  
Filed : February 27, 2002 (herewith)  
Page 6

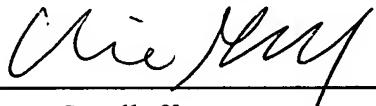
Conclusion

Applicant requests examination of the claims as amended. Should there be any minor matters preventing such examination, applicant requests that the Examiner contact the attorney indicated below.

Respectfully submitted,

AMSTER, ROTHSTEIN & EBENSTEIN  
Attorneys for Applicant  
90 Park Avenue  
New York, New York 10016  
(212) 697-5995

Dated: New York, New York  
February 27, 2002

By:   
Elie H. Gendloff  
Registration No.: 44,704

Appendix

Marked up amendments

Continuation of U.S. Patent Application 09/245,169, Filed February 27, 2002  
Added material is underlined; deleted material is bracketed.

In the Specification:

On page 14, the paragraph on lines 7-11 has been amended as follows:

Figure 5. Figure 5 depicts the DNA sequence of a PDE1C cDNA (SEQ ID NO:1) that confirms that PDE1C is expressed in pancreatic islet  $\beta$ -cells. Reverse transcriptase polymerase chain reaction was used to amplify and clone a fragment of the PDE1C mRNA common to all known PDE1C isozymes.

On page 28, the two paragraphs on lines 10-25 have been amended as follows:

--Reverse transcriptase polymerase chain reaction (RT-PCR) analysis. RT-PCR analysis was performed on 5  $\mu$ g of RNA prepared from  $\beta$ TC3 cells using Trizol (Gibco-BRL). Controls lacking reverse transcriptase were included in the reactions. To determine expression of PDE1C the following oligonucleotides were used: for RT - oligo dT; and for PCR amplification - JWPDE1C-5' -ACAGGGCAGAGGAGATCAAGTTT (SEQ ID NO:2); and JWPDE1C-3 5'-CTTTTCGCCTGCCTTTCTCCTT (SEQ ID NO:3). The 408 bp PCR product was cloned and its DNA sequence was determined.

The following oligonucleotides were used for PCR amplification to determine the expression of PDE4A: JWPDE4A-5 5'-AGCCATGGAACAGTCAAAGGTCAA (SEQ ID NO:4); and JWPDE4A-3 5'-TCAGGAGGGCCAGGAGTCGT (SEQ ID NO:5); and to determine the expression of PDE4D: JWPDE4D-5 5'-GAGGGCCGGCAGGGACAGAC (SEQ ID NO:6); and JWPDE4D-3 5'-GGGGGTGGGGTGGGTGAGAGG (SEQ ID NO:7). Amplification products 436 AND 470 bp long were obtained for PDE4A and D, respectively.--